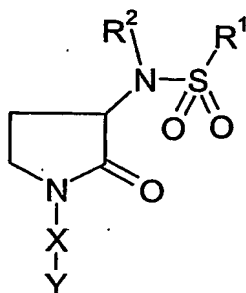


Claims

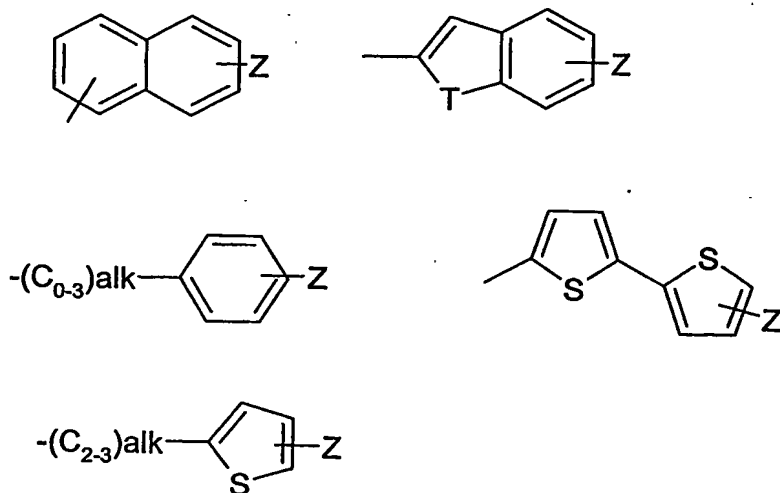
1. A compound of formula (I):



(I)

wherein:

R¹ represents a group selected from:



each ring of which optionally contains a further heteroatom N,

10 Z represents an optional substituent halogen,

alk represents alkylene or alkenylene,

T represents S, O or NH;

R² represents hydrogen, -C₁₋₆alkyl, -C₁₋₃alkylCONRᵃRᵇ, -C₁₋₃alkylCO₂C₁₋₄alkyl, -CO₂C₁-
15 4alkyl or -C₁₋₃alkylCO₂H;

Rᵃ and Rᵇ independently represent hydrogen, -C₁₋₆alkyl, or together with the N atom to which they are bonded form a 5-, 6- or 7- membered non-aromatic heterocyclic ring

optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C_{1-4} alkyl, and optionally the S heteroatom is substituted by O, i.e. represents $S(O)_n$;

5 n represents 0-2;

X represents phenyl or a 5- or 6- membered aromatic heterocyclic group containing at least one heteroatom selected from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, $-C_{1-4}$ alkyl, $-C_{2-4}$ alkenyl, $-CN$, $-CF_3$, $-NR^aR^b$, $-C_{0-4}$ alkylOR^e,
 10 $-C(O)R^f$ and $-C(O)NR^aR^b$;

R^e represents hydrogen or $-C_{1-6}$ alkyl;

R^f represents $-C_{1-6}$ alkyl;

15

Y represents a group $-C(R^x)(R^z)C_{0-2}$ alkylNR^cR^d;

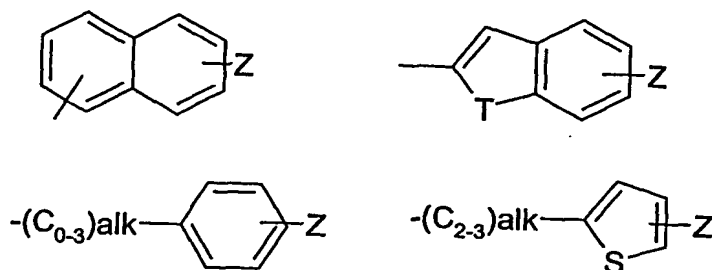
R^x represents C_{1-4} alkyl optionally substituted by halogen;

20 R^z represents hydrogen or C_{1-4} alkyl optionally substituted by halogen;

R^c and R^d independently represent hydrogen, $-C_{1-6}$ alkyl, $-C_{1-4}$ alkylOH, or together with the N atom to which they are bonded form a 4-, 5-, 6- or 7- membered non-aromatic heterocyclic ring, the 5-, 6- or 7- membered non-aromatic heterocyclic ring optionally
 25 containing an additional heteroatom selected from O, N or S, optionally substituted by C_{1-4} alkyl;
 and/or pharmaceutically acceptable derivative thereof.

2. A compound according to claim 1 wherein R¹ represents a group selected from:

30



each ring of which optionally contains a further heteroatom N,
 Z represents an optional substituent halogen,

alk represents alkylene or alkenylene,

T represents S, O or NH

and/or pharmaceutically acceptable derivative thereof.

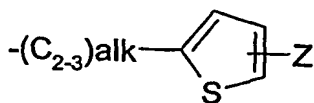
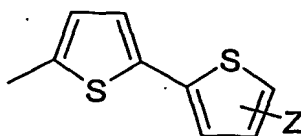
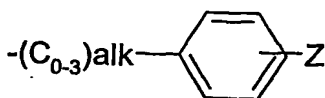
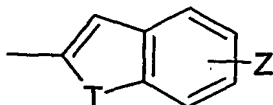
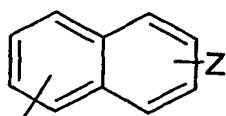
5 3. A compound according to claim 1 or claim 2 wherein R^2 represents hydrogen and/or pharmaceutically acceptable derivative thereof.

4. A compound according to any one of claims 1-3 wherein X represents phenyl or a 5- or 6- membered aromatic heterocyclic group containing at least one heteroatom selected
10 from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, $-C_{1-4}$ alkyl and $-NR^aR^b$ and/or pharmaceutically acceptable derivative thereof.

5. A compound according to any one of claims 1-4 wherein Y represents a group $-C(R^*)(R^z)NR^cR^d$ and/or pharmaceutically acceptable derivative thereof.

15

6. A compound according to claim 1 wherein R^1 represents a group selected from:



each ring of which optionally contains a further heteroatom N,

Z represents an optional substituent halogen,

alk represents alkylene or alkenylene,

20 T represents S, O or NH;

R^2 represents hydrogen, $-C_{1-6}$ alkyl, $-C_{1-3}$ alkylCONR^aR^b, $-C_{1-3}$ alkylCO₂C₁₋₄alkyl, $-CO_2$ C₁₋₄alkyl or $-C_{1-3}$ alkylCO₂H;

25 R^a and R^b independently represent hydrogen, $-C_{1-6}$ alkyl, or together with the N atom to which they are bonded form a 5-, 6- or 7- membered non-aromatic heterocyclic ring

optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl, and optionally the S heteroatom is substituted by O, i.e. represents S(O)_n;

- 5 X represents phenyl or a 5- or 6- membered aromatic heterocyclic group containing at least one heteroatom selected from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, -C₁₋₄alkyl, -C₂₋₄alkenyl, -CN, -CF₃, -NR^aR^b, -C₀₋₄alkylOR^e, -C(O)R^f and -C(O)NR^aR^b;

- 10 R^e represents hydrogen or -C₁₋₆alkyl;

R^f represents -C₁₋₆alkyl;

Y represents a group -C(R^x)(R^z)C₀₋₂alkylNR^cR^d;

15

R^x represents C₁₋₄alkyl optionally substituted by halogen (e.g. CF₃, -CH₂CF₃);

R^z represents hydrogen or C₁₋₄alkyl optionally substituted by halogen (e.g. CF₃, -CH₂CF₃);

- 20 R^c and R^d independently represent hydrogen, -C₁₋₆alkyl, -C₁₋₄alkylOH, or together with the N atom to which they are bonded form a 5- or 6- membered non-aromatic heterocyclic ring optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl;
and pharmaceutically acceptable derivatives thereof.

25

7. A compound according to claim 1 selected from:

(E)-2-(5-Chloro-2-thienyl)-N-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethanesulfonamide;

(E)-2-(5-Chloro-2-thienyl)-N-(1-{2-fluoro-4-[1-(4-morpholinyl)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)ethanesulfonamide;

30

(E)-2-(5-Chloro-2-thienyl)-N-[1-(2-fluoro-4-{1-[(2-hydroxyethyl)(methyl)amino]ethyl}phenyl)-2-oxo-3-pyrrolidinyl]ethanesulfonamide;

(E)-N-{1-[4-(1-Aminoethyl)-2-fluorophenyl]-2-oxo-3-pyrrolidinyl}-2-(5-chloro-2-thienyl)ethanesulfonamide;

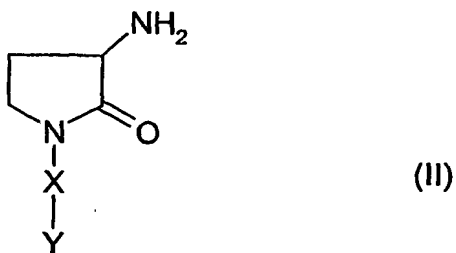
35 6-Chloro-N-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;

(E)-2-(5-Chloro-2-thienyl)-N-((3S)-1-{4-[(1S)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethanesulfonamide;

- 6-Chloro-*N*-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 5 (E)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 6-Chloro-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 10 (E)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (E)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 15 (E)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[1-(dimethylamino)propyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (E)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[1-(dimethylamino)-2-methylpropyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (E)-2-(5-Chloro-2-thienyl)-*N*-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 20 6-Chloro-*N*-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 25 6-Chloro-*N*-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-(1-{4-[1-(ethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-[1-(4-{1-[ethyl(methyl)amino]ethyl}-2-fluorophenyl)-2-oxo-3-pyrrolidinyl]-2-naphthalenesulfonamide;
- 30 6-Chloro-*N*-[1-(2-fluoro-4-{1-[(1-methylethyl)amino]ethyl}phenyl)-2-oxo-3-pyrrolidinyl]-2-naphthalenesulfonamide;
- 6-Chloro-*N*-[1-(2-fluoro-4-{1-[methyl(1-methylethyl)amino]ethyl}phenyl)-2-oxo-3-pyrrolidinyl]-2-naphthalenesulfonamide;
- 35 *N*-(1-{4-[1-(1-Azetidinyl)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-6-chloro-2-naphthalenesulfonamide;

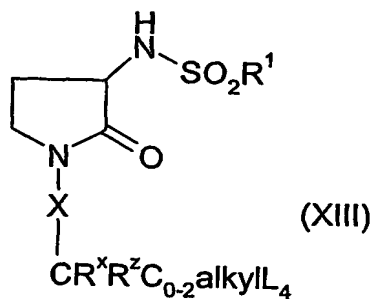
- 6-Chloro-*N*-(1-{2-fluoro-4-[1-(1-pyrrolidinyl)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-*N*-(1-{2-fluoro-4-[1-(1-piperidinyl)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 5 5'-Chloro-*N*-((3*S*)-1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2,2'-bithiophene-5-sulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 10 6-Chloro-*N*-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 6-Chloro-*N*-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 15 (1*E*)-2-(5-Chloro-2-thienyl)-*N*-(1-{4-[1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)-1-propene-1-sulfonamide;
- 6-Chloro-*N*-(1-{4-[1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- and/or pharmaceutically acceptable derivative thereof.
- 20
8. A compound according to any one of claims 1-7 and/or pharmaceutically acceptable derivative thereof for use in therapy.
9. A pharmaceutical composition comprising a compound according to any one of claims
- 25 1-7 and/or pharmaceutically acceptable derivative thereof together with at least one pharmaceutical carrier and/or excipient.
10. Use of a compound according to any one of claims 1-7 and/or pharmaceutically acceptable derivative thereof for the manufacture of a medicament for the treatment of a
- 30 patient suffering from a condition susceptible to amelioration by a Factor Xa inhibitor.
11. A method of treating a patient suffering from a condition susceptible to amelioration by a Factor Xa inhibitor comprising administering a therapeutically effective amount of a compound according to any one of claims 1-7 and/or pharmaceutically acceptable
- 35 derivative thereof.
12. A process for preparing a compound of formula (I) which comprises:

(a) reacting compounds of formula (II) or an acid addition salt thereof with compounds of formula (III) where V is a suitable leaving group:



OR:

(b) by reacting compounds of formula (XIII) with HNR^cR^d :



OR:

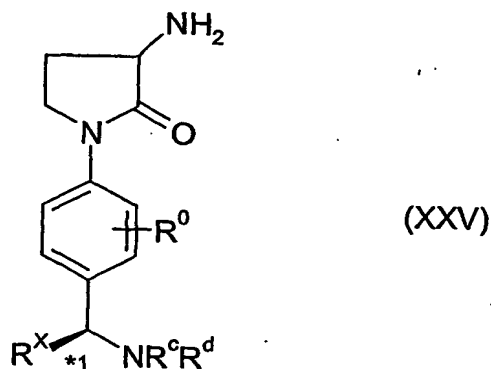
(c) by reacting compounds of formula (I) where R^2 is hydrogen with compounds of formula (XVII):



wherein R^2 is $-\text{C}_{1-6}\text{alkyl}$, $-\text{C}_{1-3}\text{alkylCONR}^a\text{R}^b$, $-\text{C}_{1-3}\text{alkylCO}_2\text{C}_{1-4}\text{alkyl}$, or $-\text{CO}_2\text{C}_{1-4}\text{alkyl}$ and T is a suitable leaving group, optionally followed by removal of the alkyl protecting group where appropriate;

OR:

(d) by reacting a compound of formula (XXV) where X represents phenyl, Y represents $\text{CH}(\text{R}^x)\text{NR}^c\text{R}^d$, R^c and R^d each represent the same C_{1-6} alkyl substituent and R^0 represents 0-2 optional substituents on the phenyl ring selected from: halogen, -C_{1-4} alkyl, -C_{2-4} alkenyl, -CN , -CF_3 , $\text{-NR}^a\text{R}^b$, -C_{0-4} alkylOR^e, -C(O)R^f and $\text{C(O)NR}^a\text{R}^b$ and/or an acid addition salt thereof:

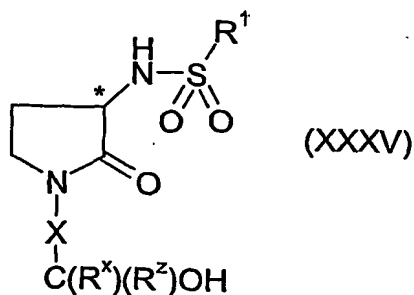


with a compound of formula (III) where V is a suitable leaving group:



OR:

(e) treatment of compounds of formula (XXXV) where Y represents $\text{-C(R}^x)(\text{R}^z)\text{NR}^c\text{R}^d$ and R^x and R^z both represent C_{1-4} alkyl and R^2 represents hydrogen:

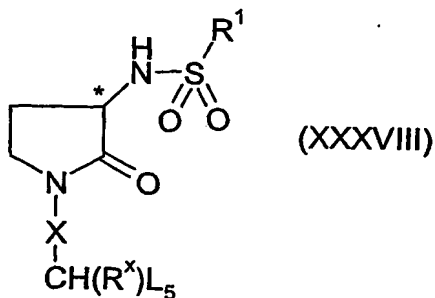


with hydrogen chloride in the presence of zinc chloride, followed by reaction with HNR^cR^d ;

OR:

(f) by reacting compounds of formula (XXXVIII) where Y represents $-C(R^x)NR^cR^d$, R^x represents C_{1-4} alkyl and R^c and R^d independently represent hydrogen, C_{1-6} alkyl, or together with the N atom to which they are bonded form a 4-, 5-, 6- or 7- membered non-aromatic heterocyclic ring and L_5 is a suitable leaving group:

5



with HNR^cR^d .